October 14, 2005

CLAIMS

- 1. A attenuated myxomavirus or components thereof for use as monoparamunity inducer.
- 2. The monoparamunity inducer as claimed in claim 1, characterized in that the attenuated myxomavirus strain has a modification in the form of an addition, substitution or deletion in one or more gene segments coding for the production of cytokine receptors, with the receptor properties of the cytokine receptor being lost through the modification.
- 3. The monoparamunity inducer as claimed in claim 2, characterized in that the cytokine receptors are selected from the group of receptors for interferons (IFN), interleukins (IL) and tumor necrosis factor (TNF).
- 4. The monoparamunity inducer as claimed in claim 3, characterized in that the cytokine receptors are selected from the group of receptors for IFN α -R, IFN γ -R, TNF-R, IL-1-R, IL-2-R, IL-6-R and IL-12-R.
- 5. The monoparamunity inducer as claimed in any of claims 1 to 4, characterized in that the viral components include viral envelopes or aberrant forms of viral envelopes of an attenuated myxomavirus strain.
- 6. The monoparamunity inducer as claimed in any of claims 1 to 5, characterized in that the myxomavirus strain is selected from the group consisting of M2, M7, Lausanne, Aust/Uriarra/Verg-86/1.
- 7. The monoparamunity inducer as claimed in claim 6,

characterized in that the myxomavirus strain is the strain M-2 with the deposit number ECACC 03121801.

- 8. The monoparamunity inducer as claimed in any of claims 1 to 7, characterized in that the myxomavirus strain has been attenuated in cell cultures.
- 9. The monoparamunity inducer as claimed in claim 8, characterized in that the cell cultures comprise Vero monkey kidney cells and/or AVIVER cells.
- 10. The monoparamunity inducer as claimed in any of claims 1 to 9, characterized in that the attenuated myxomaviruses are inactivated.
- 11. The monoparamunity inducer as claimed in claim 10, characterized in that the attenuated myxomaviruses have been inactivated with beta-propiolactone.
- 12. The monoparamunity inducer as claimed in claim 11, characterized in that the concentration of beta-propiolactone is 0.01-1%.
- 13. The monoparamunity inducer as claimed in claim 12, characterized in that the concentration of beta-propiolactone is 0.05%.
- 14. The monoparamunity inducer as claimed in any of claims 1 to 13, characterized in that the myxomaviruses have been lyophilized.
- 15. The use of the monoparamunity inducer as claimed in any of claims 1 to 14 for producing a pharmaceutical composition for activating the paraspecific immune system in humans and animals.
- 16. The use of the monoparamunity inducer as claimed in any of claims 1 to 14 for producing a pharmaceutical

composition for the parenteral treatment and/or for the dysfunctions of the immune prophylaxis of immunosuppression, immunodeficiency disorders, dysfunctions of homeostasis between the hormonal, circulatory, metabolic and nervous systems, threatened neonatal infection, neoplastic diseases, diseases, bacterial diseases, therapy-resistant infectious factor diseases, viral and bacterial mixed infections, chronic manifestations οf infectious processes, liver disorders of various etiologies, chronic skin disorders, herpetic diseases, chronic hepatitides, influenzal infections, endotoxin damage.

- 17. The use as claimed in any of claims 15 to 16, where the treatment takes place by local administration of the pharmaceutical composition via the skin or mucous membranes of patient.
- 18. A pharmaceutical composition including a monoparamunity inducer as claimed in any of claims 1 to 14 and a pharmaceutically acceptable carrier.
- 19. The pharmaceutical composition as claimed in claim 18, for local or parenteral administration.
- 20. The pharmaceutical composition as claimed in claim 18, where the composition is in the form of buccal and suckable tablets.
- 21. A method for producing a monoparamunity inducer based on an attenuated rabbit myxomavirus strain, comprising the steps:
- isolation of myxomaviruses from infected tissue of a rabbit typically suffering from generalized myxomatosis;
- adaptation of the virus to a permissive cell system;
 - passaging of the isolated viruses in a Vero and

binary cell culture until attenuation of the virus is achieved.

- 22. The method as claimed in claim 21, where the isolated viruses are inoculated onto the chorioallantoic membrane (CAM) of incubated chicken eggs for the adaptation.
- 23. The method as claimed in claim 22, where the virus is replicated over 2 to 6 passages, preferably over 3 passages, on the CAM.
- 24. The method as claimed in claim 21, where the myxomaviruses present in the infected tissue are isolated by replication in a permissive cell system.
- 25. The method as claimed in claim 24, characterized in that the virus is isolated by culturing on the chorioallantoic membrane (CAM) of incubated chicken eggs.
- 26. The method as claimed in claim 25, characterized in that the virus is subsequently adapted to the CAM over further passages, preferably over 2 further passages.
- 27. The method as claimed in any of claims 21 to 26, characterized in that the attenuation of the myxomaviruses takes place over 80 to 150 passages, preferably over 120 passages, in Vero cell cultures.
- 28. The method as claimed in any one of claims 21 to 27, characterized in that the virus is passaged in an AVIVER cell culture.
- 29. The method as claimed in claim 28, characterized in that the passaging takes place over 10 to 50 passages, preferably over 25 passages.

- 30. The method as claimed in any of claims 21 to 29, where the attenuated myxomaviruses are additionally replicated over further attenuation passages.
- 31. The method as claimed in claim 30, where the replication takes place in Vero monkey kidney cells.
- 32. The method as claimed in claim 31, characterized in that the further growth takes place in Vero cells over 100 to 200 passages.
- 33. The method as claimed in any of claims 30 to 32, characterized in that the viral harvests of the cell culture have an infectious titer of from 10^5 to $10^{7.5}$, preferably of at least $10^{6.5}$, $TCID_{50}/ml$.
- 34. The method as claimed in any of claims 21 to 33, where the attenuated myxomaviruses are additionally inactivated.
- 35. The method as claimed in claim 34, where the attenuated myxomaviruses are treated with beta-propiolactone for the inactivation.
- 36. The method as claimed in claim 35, characterized in that the concentration of beta-propiolactone is 0.01-1%.
- 37. The method as claimed in claim 36, characterized in that the concentration of beta-propiolactone is 0.05%.
- 38. The method as claimed in any of claims 21 to 37, including the steps
- isolation of myxomaviruses from the infected tissue of the rabbit suffering from myxomatosis by replication on the CAM of incubated chicken eggs and

subsequent

- adaptation of the isolated myxomavirus to the CAM over a further 2 passages
 - attenuation of the isolated viruses by
 - passaging in Vero cell cultures, preferably over 120 passages,
 - transfer of the viruses into the binary AVIVER cell culture with further attenuation of the virus, preferably over 24 intermediate passages, in this cell culture,
 - subsequent transfer of the virus back onto
 Vero monkey kidney cells, and
- replication of the attenuated myxomaviruses by further attenuation passages in the Vero cells, preferably over about 150 passages.
- 39. An attenuated myxomavirus obtainable by the method as claimed in any of claims 21 to 38.
- 40. The myxomavirus as claimed in claim 39, deposited at the European Collection of Cell Cultures (ECACC) under the deposit number 03121801.
- 41. The myxomavirus as claimed in claim 39 or 40, characterized in that the myxomavirus is genetically modified.
- claimed in claim The myxomavirus as 41, characterized in that the myxomavirus have modification in the form of an addition, substitution or deletion in one or more gene segments coding for the production of cytokine receptors, with the receptor properties of the cytokine receptor being lost through the modifications.
- 43. The myxomavirus as claimed in claim 42, characterized in that the cytokine receptors are

selected from the group of receptors for interferons (IFN), interleukins (IL) and tumor necrosis factor (TNF).

- 44. The myxomavirus as claimed in claim 43, characterized in that the cytokine receptors are selected from the group of receptors for IFN α -R, IFN γ -R, TNF-R, IL-1-R, IL-2-R, IL-6-R and IL-12-R.
- 45. An attenuated myxomavirus strain M2 with the deposit number ECACC 03121801.